

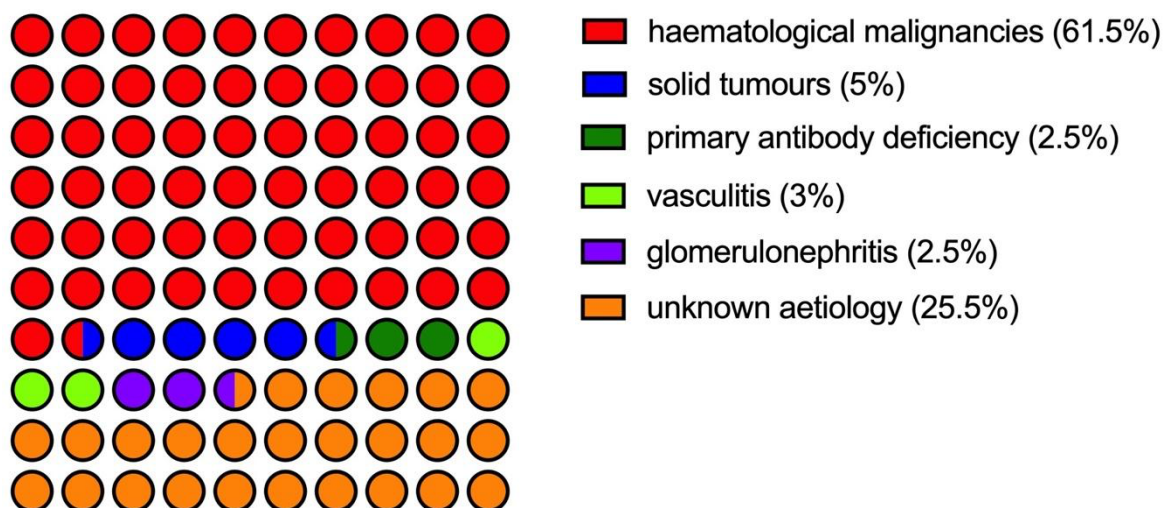
SUPPLEMENTARY MATERIAL

Microbiological evidences	HYPO (n=39)	no-HYPO (n=335)
<i>Staphylococcus aureus</i>	5	3
<i>Streptococcus pneumoniae</i>	5	3
<i>Enterococcus faecium</i>	2	2
<i>Escherichia coli</i>	2	2
<i>Pseudomonas aeruginosa</i>	2	1
<i>Cytomegalovirus</i>	3	0
<i>Aspergillus fumigatus</i>	2	0
<i>Clostridium difficile</i>	0	1
<i>Klebsiella pneumoniae</i>	3	1

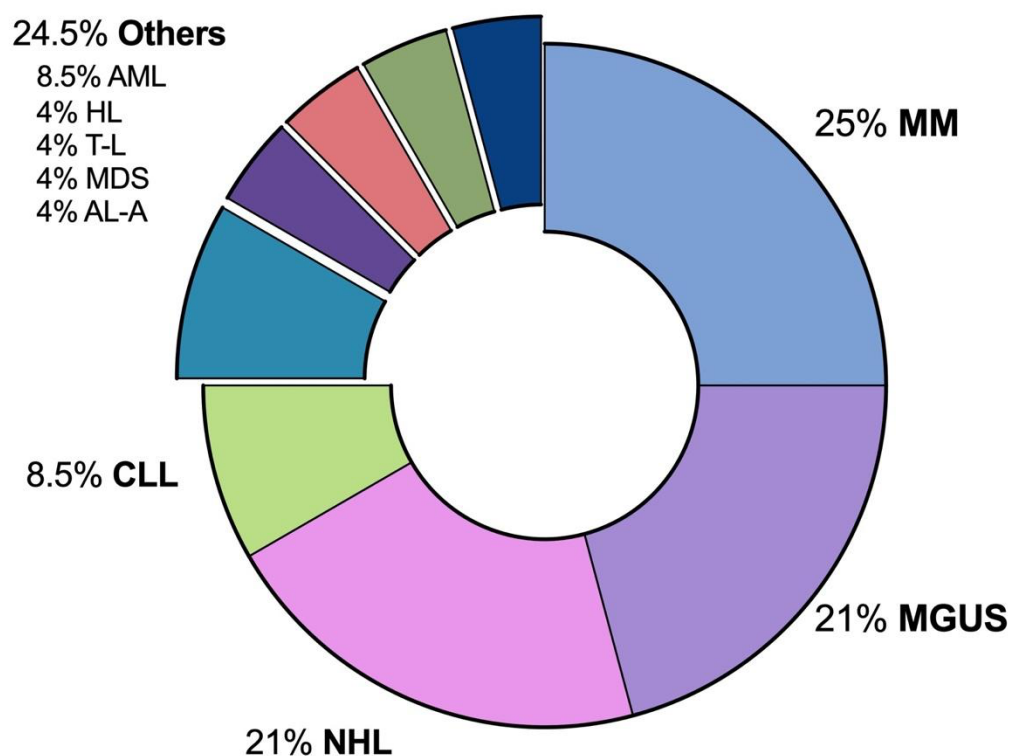
Supplementary Table 1.

Different pathogens in microbiological samples (blood cultures, urine cultures and swabs) between HYPO and no-HYPO groups in our COVID-19 cohort.

HYPO = hypogammaglobulinemia.

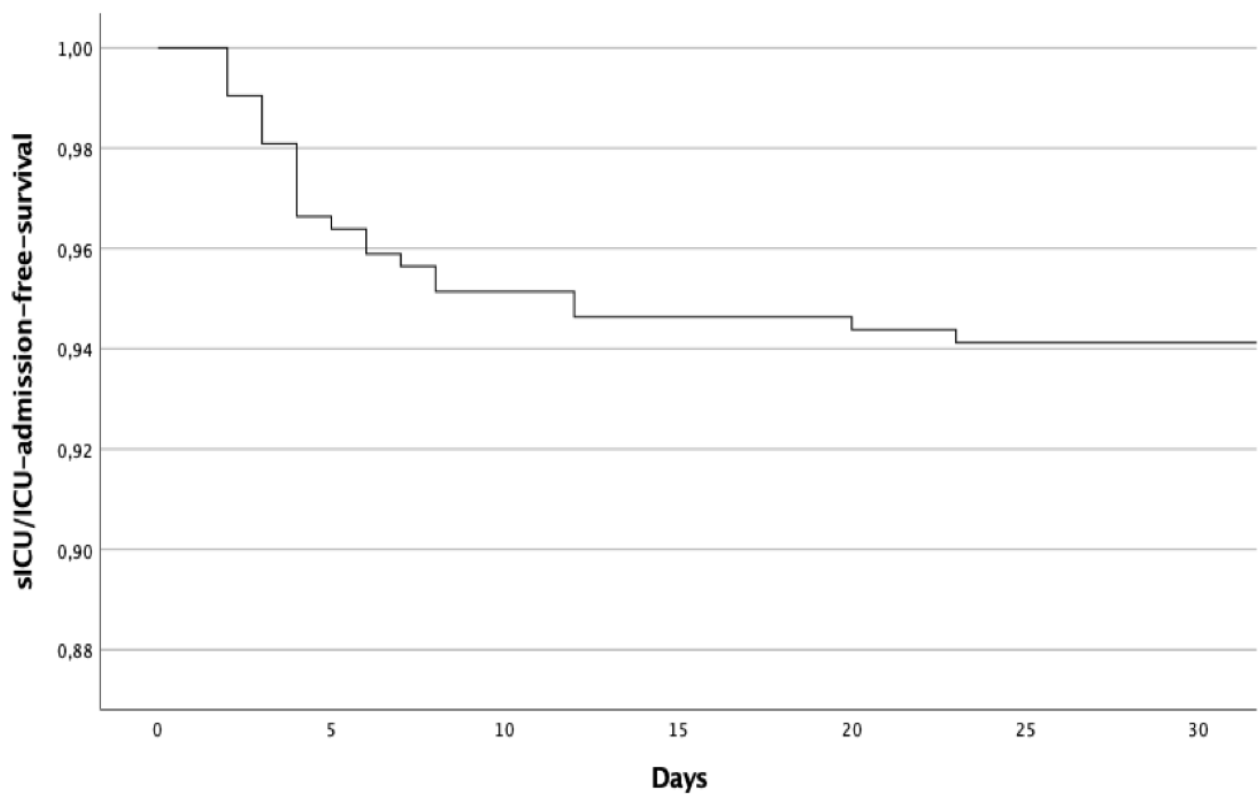


Supplementary figure 1. Different aetiologies of hypogammaglobulinemia in our COVID-19 cohort are expressed as percentage of the total (HYPO group, n = 39 cases).

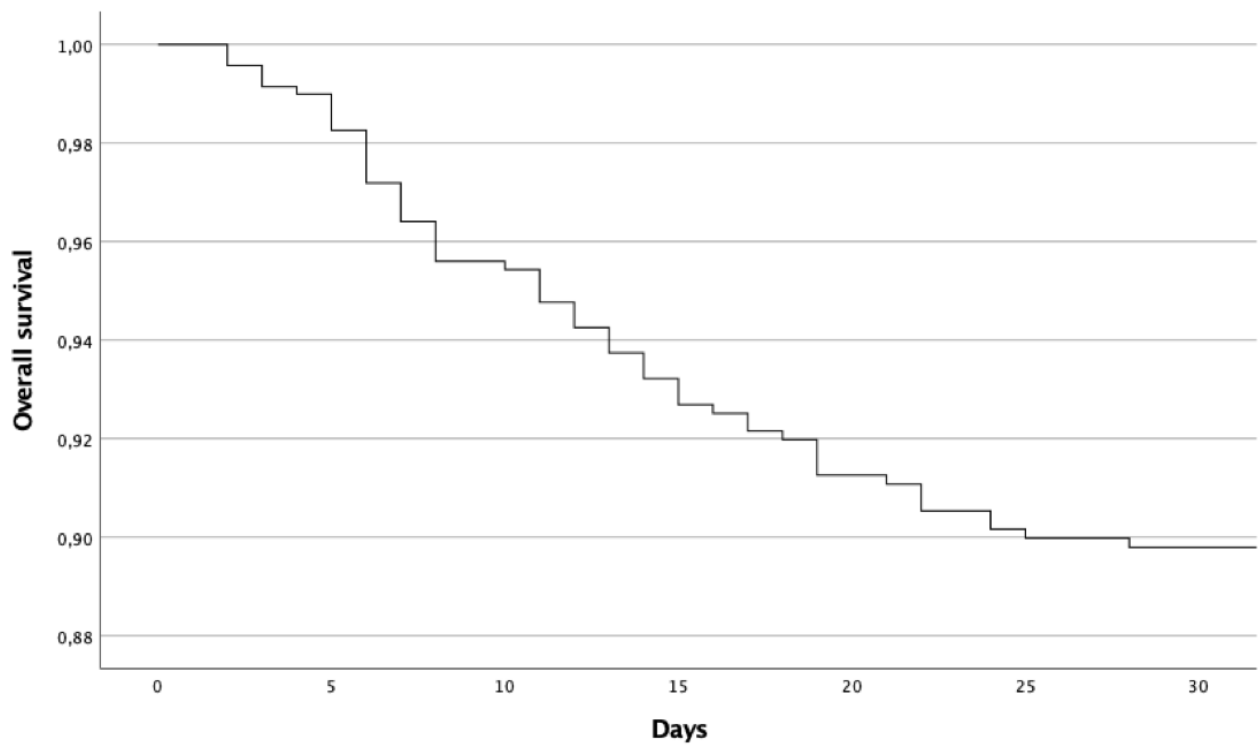


Supplementary Figure 2. Different causes of hypogammaglobulinemia in the haematological subgroup of hypogammaglobulinemia (n = 24, 61.5% of HYPO group).

MGUS = monoclonal gammopathy of undetermined significance, MM = multiple myeloma, NHL = non-Hodgkin's lymphoma, CLL = chronic lymphocytic leukaemia, AML = acute myeloid leukaemia, HL = Hodgkin's lymphoma, T-L = T-cell lymphoma, MDS = myelodysplastic syndrome, AL-A = AL amyloidosis.

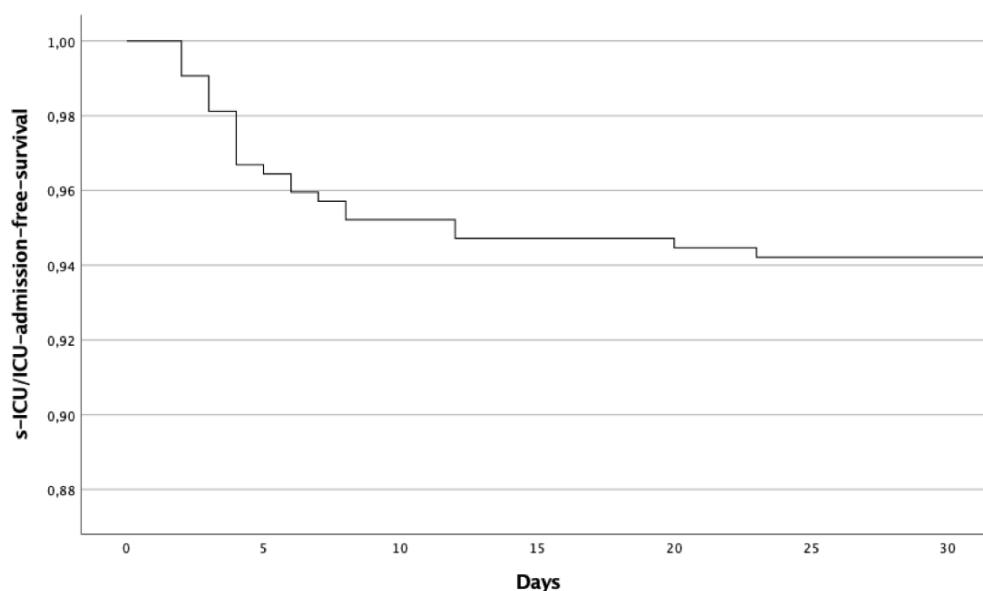


Supplementary Figure 3 A. Survival analysis of sICU/ICU admission within 30 day adjusted for HYPO, age, sex, cancer and steroid therapy. Days are calculated since the admission to the hospital. Cox Regression HR adjusted: 3.73 (1.75-7.91), $p < 0.001$.



Supplementary Figure 3 B. Survival analysis of mortality within 30 day since admission adjusted for HYPO, age, sex and cancer and steroid therapy. Days are calculated since the admission to the hospital. . Cox Regression HR adjusted: 1.42 (0.62-3.23), $p = 0.404$.

Haematological malignancies were differently represented between HYPO and non-HYPO patients, as evidenced by the binomial regression analysis ($p < 0.001$, OR 4.69. CI95% 2.14-10.31). However, when replacing cancer with hematological malignancies in the multivariate regression analyses, the significance of the model and the impact of hypogammaglobulinemia did not change. When considering the outcome “ICU admission” we found $p = 0.002$ for hypogammaglobulinemia, with OR 4.75 CI95% 1.80-12.52, whereas $p = 0.166$ for hematological malignancies, with OR 2.08 CI95% 0.74-5.85. In case of Severity score ≥ 5 we found $p < 0.001$ for hypogammaglobulinemia, with OR 12.78 CI95% 4.52-36.14, whereas $p = 0.784$ for hematological malignancies, with OR 0.86 CI95% 0.29-2.54. As a second control, we adjusted for hematological malignancies, age, sex and hypogammaglobulinemia the Cox proportional hazard regression for sICU/ICU admission, again finding a consistency of hypogammaglobulinemia with a HR of 2.86 (1.15-7.13), $p = 0.024$ (see Supplementary Figure 4 below).



Supplementary Figure 4. Adjust survival analysis of sICU/ICU admission within 30 day for HYPO, age, sex and hematological malignancies. Days are calculated since the admission to the hospital. Cox Regression HR: 2.86 (1.15-7.13), $p = 0.024$.

	ICU (n=15)	no-ICU (n=24)	<i>p</i> *
Comorbidities n (%)			
<i>Cancer</i>	10 (66.7%)	16 (66.7%)	1
<i>Active cancer treatment</i>	7 (70%)	11 (68.8%)	0.946
<i>Bronchiectasis</i>	6 (40%)	3 (12.5%)	0.047
<i>Arterial hypertension</i>	8 (53.3%)	11 (45.8%)	0.648
<i>Dyslipidemia</i>	5 (33.3%)	7 (29.2%)	0.784
<i>Diabetes mellitus</i>	2 (13.3%)	4 (16.7%)	0.779
<i>Ischemic heart disease</i>	2 (13.3%)	5 (20.8%)	0.553
<i>Obesity</i>	1 (6.7%)	4 (16.7%)	0.363
<i>Previous corticosteroid therapy</i>	8 (53.3%)	8 (33.3%)	0.217

Supplementary Table 2.

Comparison of comorbidities between ICU vs. no-ICU among the HYPO cohort.

*Chi-squared test.